

Review Article

The use of psychedelic agents in the treatment of alcohol and tobacco dependence: An integrative review

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Objective: to analyze studies that address the use of psychedelic agents in the treatment of alcohol and tobacco dependence in the national and international contexts. Method: an integrative review in Literatura Latino Americana e do Caribe em Ciências da Saúde (LILACS), Medical Literature Analysis and Retrieval System Online (Medline), Scientific Electronic Library Online (SciELO), EBSCO and PubMed, using the following descriptors: "Alucinógenos", "Tratamento", "Dependência Química", "Alcoolismo", "Tabagismo", "Hallucinogens", "Treatment", "Chemical Dependence", "Alcoholism" and "Tobacco Use Disorder". **Results**: the final sample totaled 14 articles that addressed the use of psychedelic agents in the treatment of alcohol and tobacco dependence in the national and international contexts. Conclusion: the use of psychedelic agents in the treatment of alcohol and tobacco dependence may not be conclusive in the treatment of chemical dependence as a single therapy; however, its use can be an effective innovation to the current treatment models, so that this includes the patient's psychedelic experience and its relationship with treatment.

Descriptors: Hallucinogens; Treatment; Chemical Dependence; Alcoholism; Tobacco Use Disorder.

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O uso de agentes psicodélicos no tratamento da dependência de álcool e tabaco: revisão integrativa

Objetivo: analisar os estudos que abordam o uso de agentes psicodélicos no tratamento da dependência de álcool e tabaco no contexto nacional e internacional. **Método**: revisão integrativa na Literatura Latino-Americana e do Caribe em Ciências da Saúde (LILACS), *Medical Literature Analysis and Retrievel System Online* (Medline), *Scientific Eletronic Library Online* (SciELO), EBSCO e PubMed, utilizando-se os descritores: "Alucinógenos", "Dependência Química", "Tratamento", "Alcoolismo", "Tabagismo", "Hallucinogens", "Chemical Dependence", "Treatment", "Alcoholism", "Tobacco Use Disorder". **Resultados**: a amostra final totalizou 14 artigos, que abordaram o uso de agentes psicodélicos no tratamento de dependência de álcool e tabaco, no contexto nacional e internacional. **Conclusão**: o uso de agentes psicodélicos no tratamento da dependência química como terapêutica única, entretanto, sua utilização pode ser uma inovação eficaz para os modelos atuais de tratamentos, de modo que engloba a experiência psicodélica do paciente e a relação desta com o tratamento.

Descritores: Alucinógenos; Terapêutica; Transtornos Relacionados ao Uso de Substâncias; Terapia; Alcoolismo; Tabagismo.

El uso de agentes psicodélicos en el tratamiento de la dependencia de alcohol y tabaco: revisión integradora

Objetivo: analizar estudios que aborden el uso de agentes psicodélicos en el tratamiento de la adicción al alcohol y al tabaco en el contexto nacional e internacional. **Método**: revisión integradora en Literatura Latinoamericana y del Caribe en Ciencias de la Salud (LILACS), *Medical Literature Analysis and Retrievel System Online* (Medline), *Scientific Eletronic Library Online* (SciELO), EBSCO y PubMed, utilizando los siguientes descriptores: "Alucinógenos", "Tratamento", "Dependência Química", "Alcoolismo", "Tabagismo", "Hallucinogens", "Treatment", "Chemical Dependence", "Alcoholism" y "Tobacco Use Disorder". **Resultados**: la muestra final consideró 14 artículos que abordaron el uso de agentes psicodélicos en el tratamiento de la adicción al alcohol y al tabaco en el contexto nacional e internacional. **Conclusión**: el uso de agentes psicodélicos en el tratamiento de la dependencia química como terapia única; sin embargo, su uso puede implicar una innovación eficaz a los modelos de tratamiento actuales, dado que incluye la experiencia psicodélica del paciente y la relación de la misma con el tratamiento.

Descriptores: Alucinógenos; Tratamiento; Dependencia Química; Alcoholismo; Tabaquismo.

Introduction

The use of psychoactive substances accompanies humanity's history and culture. In the search for pleasure, and also to heal distresses, several substances have been discovered, whether natural or artificial. However, there is a set of harmful effects that are caused when there is chemical dependence on these substances⁽¹⁾.

Globally, alcohol dependence is the most prevalent in the world, and harmful alcohol use is responsible for 3 million deaths a year worldwide, in addition to being a factor for more than 200 injuries and diseases⁽²⁾. In Brazil, the 2017 National Survey on Drug Use by the Brazilian Population exposes abuse and dependence on numerous substances, showing alarming data, mainly regarding the use of licit drugs such as alcohol and tobacco⁽³⁾. Tobacco kills more than seven million people every year; even though its consumption has decreased significantly since 2000, it is still insufficient to avoid the consequences of using this substance, such as cardiovascular diseases and other chronic non-communicable diseases, including 44% of all deaths⁽⁴⁾. If the number of consumers is high, the sample space increases, which consequently covers people pre-disposed to the development of a possible chemical dependence.

As it is a public health problem, the Brazilian State spends considerable amounts of money for the treatment of citizens who have chemical dependence⁽⁵⁾. Initially, dependent use of tobacco – which is one of the most prevalent in Brazil – harms the State economically by approximately 60 billion reais a year due to medical expenses for treatment, and carries with it nearly 160,000 deaths that could be avoided⁽⁶⁾. Smoking causes dependence, initially on nicotine, which stimulates the brain to release various neurotransmitters, inducing the body to a sense of pleasure⁽⁷⁾.

Together, according to a survey by researchers from the Oswaldo Cruz Foundation (FIOCRUZ), the Federal University of Rio de Janeiro (Universidade Federal do Rio de Janeiro, UFRJ), the State University of Rio de Janeiro (Univerdade Estadual do Rio de Janeiro, UERJ) and the Health Surveillance Secretariat of the Ministry of Health, the financial impact to treat diseases related to alcoholism by the Unified Health System (Sistema Único de Saúde, SUS) is estimated at around 8.2 million reais a year, and reaches nearly 10% of the Brazilian population⁽⁸⁾. Alcoholism tends to be attributed to environmental factors, such as: 1) Cultural development; 2) Economy; 4) Alcohol availability. Even if there is not only one dominant factor, the greater the vulnerability in which the individuals are in, the more likely they are to develop problems related to alcohol and its use⁽⁹⁾; therefore, it is remarkable that chemical dependence, especially on alcohol and tobacco, is a relevant condition for the government, in social and economic terms.

The treatment of alcoholism is based on several aspects, which are determined according to the severity of dependence, but which are distributed from Primary Health Care (PHC) to specialty centers, ranging from continuous follow-up therapies to drugs that control the alcohol withdrawal syndrome⁽¹⁰⁾. On the other hand, treatment of the smoking habit is based on nicotine replacement therapy, aiming to minimize the symptoms of the withdrawal syndrome, as well as on the use of the bupropion hydrochloride drug⁽¹¹⁾.

Due to the search for new treatments, the scientific community added to the group of therapeutic alternatives the use of psychedelic agents for the treatment of dependence on psychoactive substances⁽¹²⁾, as an example we have psilocybin, a hallucinogenic substance found in mushrooms of the *Psilocybe cubensis* species, which, after scientific studies, was discovered as with therapeutic potential for the treatment of chemical dependence⁽¹³⁾.

The scientific community has been looking at new therapeutic alternatives for alcoholics and smokers; in this sense, this research conducted a survey of bibliographic productions on the use of psychedelic agents in the treatment of alcohol and tobacco dependence in the national and international contexts. The objective was to identify and describe the treatments of the tobacco and alcohol addictions conducted with the use of psychedelic agents, aiming to assist in increasing the sampling of therapeutic alternatives.

Method

This is a study of the Integrative Literature Review type, whose methodology aims at synthesizing results obtained from a broad analysis of the literature on a given theme or question, in a systematic and orderly manner, contributing to discussions on the research methods and results⁽¹⁴⁾.

Preparation of this integrative review study followed six stages: 1) Identification of the theme and selection of the research question for the elaboration of the integrative review; 2) Establishment of criteria for inclusion and exclusion of studies/sampling; 3) Definition of the information to be extracted from the studies selected, as well as categorization of the studies; 4) Evaluation of the studies included in the integrative review; 5) Interpretation of the results; and 6) Presentation of the review and synthesis of the knowledge⁽¹⁵⁾. In addition, in order to guide the integrative review, the following guiding question was elaborated: Which is the effect of psychedelic agents on the treatment of alcohol and tobacco dependence reported in the scientific production from 2010 to 2020?

The articles were searched and selected jointly in July 2020. All the articles selected were compared with each other in order to verify repetition and agreement of the data obtained, together with the inclusion and exclusion criteria. Observational and experimental studies published in the last 10 years (from 2010 to 2020), with free access, in English, Portuguese and Spanish, and which evaluate the use of psychedelic agents in the treatment of alcohol and tobacco dependence in adults, were included. Articles published before 2010 were excluded, as well as paid articles, articles that do not present adequate reasoning, editorials, reviews, opinion articles, and other studies that do not answer the main research question.

4

The following databases were consulted: *Literatura Latino-Americana e do Caribe em Ciências da Saúde* (LILACS), Medical Literature Analysis and Retrieval System Online (Medline), Scientific Electronic Library Online (SciELO), EBSCO and PubMed. Thus, the research was able to encompass, with these databases, new articles in the national and international contexts.

The descriptors used were defined based on the Descriptors in Health Sciences, with the following terms and their combinations, separated by the Boolean operators "AND" and "OR", in Portuguese and English: : "Alucinógenos", "Dependência Química", "Tratamento", "Alcoolismo", "Tabagismo", "Hallucinogens", "Chemical Dependence", "Treatment", "Alcoholism" and "Tobacco Use Disorder", in combinations that totaled 9 different search strategies (Figure 1).

SEARCH TERMS	SCIELO	LILACS	EBSCO	PUBMED	MEDLINE
"Alucinógenos" OR "Agentes psicodélicos"	37	25	373	7	312
"Alucinógenos" OR "Agentes psicodélicos" " AND "Tratamento"	2	10	192	0	189
"Agentes psicodélicos" OR "alucinógenos" AND "tratamento" AND "dependência química" OR "adição a drogas"	1	0	152	0	189
"Hallucinogens" OR "Psychedelic Agents"	38	53	2,113	304	325
"Hallucinogens" OR "Psychedelic Agents" AND "treatment"	4	18	1,594	189	98
"Hallucinogens" OR "Psychedelic Agents" AND "treatment" AND "Chemical Dependence" OR "Drug addiction"	4	14	60	67	28
"Agentes psicodélicos" OR "alucinógenos"	37	25	373	7	312
"Agentes psicodélicos" OR "alucinógenos" AND "tratamiento"	2	5	348	0	189
"Agentes psicodélicos" OR "alucinógenos" AND "tratamiento" AND "dependência química" OR "Adicción a Drogas"	2	2	109	0	25

Figure 1 - Search terms used. Palmas, TO, Brazil, 2020

The initial search included 127 articles in Scielo, 152 in LILACS, 5,314 in EBSCO, 554 in PUBMED and 1,667 in MEDLINE, totaling 7,834 articles and, after reading their titles and abstracts, only 61 articles remained. Of these, after applying the inclusion and exclusion criteria, 14 articles remained that portrayed the use of psychedelic agents in the treatment of alcohol and tobacco dependence, in the national and international contexts.

For analysis and subsequent synthesis of the articles selected, a synoptic table was used, including the following: title, author, year, database, study objective, methodology, result, analysis and conclusion.

Results

The studies were predominantly published in the period from 2014 to 2019, with only one study published in 2011⁽¹⁶⁾. As for the origin and language of the publications, eleven articles were published in English, eight of which

came from the United States⁽¹⁷⁻²⁴⁾, one from Canada⁽²⁵⁾, one from the United Kingdom⁽²⁶⁾ and the other without identification of its locus⁽²⁷⁾; two articles in Portuguese from Brazil^(16,28), and a single article published in Spanish, in Spain⁽²⁹⁾.

Regarding the type of study, predominance of quantitative studies was identified, of which: seven were clinical trials and three were open pilot studies, in which two articles worked with the same study, where moderate (20 mg/70 kg) and high (30 mg/70 kg) doses of psilocybin were used in a smoking cessation treatment with 15 participants, during 15 weeks^(17,20), and the other study used psilocybin for the treatment of alcohol dependence, with doses of 0.3 mg/kg and 0.4 mg/kg, in different sessions, over 12 weeks, with 10 participants⁽¹⁹⁾; a double blind placebo-controlled treatment with psilocybin on alcohol use dependence, lasting 12 weeks⁽¹⁸⁾; two experimental studies conducted with mice, in one, the potential of acute administration of ayahuasca (1.76, 3.0, 17.6 and

30.0 mg/kg of DMT o.r.) was evaluated in preventing the expression of cocaine-induced behavioral sensitization, and the neurochemical repercussions of the treatment in C57BI/6⁽²⁸⁾ and, in the other study, there was administration of a non-selective 4-Acetoxi-Dimethyltryptaline (4-AcO-DMT) agonist, in Wistar rats and C57BI/6 mice, which were induced to dependence on opiates, nicotine and BNDF intra-VTA, and previously treatment virgins⁽²⁵⁾. In addition to these, a report describing the biologically verified results of tobacco abstinence in one of the pilot studies selected and related data on the subjective effects of psilocybin⁽²⁴⁾; and two online observational and cross-sectional research studies, one with a sample of 343 adults, and the other with 358^(21,23).

As for the qualitative research studies, one was of the ethnographic type⁽²⁷⁾, based on long-term fieldwork and participant observation in ayahuasca communities, and in-depth semi-structured interviews with participants who have a history of substance abuse, conducted in the context of the Chemical Youth Advanced Grant program financed by the European Research Council, which uses research methods from medical anthropology, and science and technology studies to explore the effects experienced, pragmatic regimes of the use of chemical and pharmaceutical products from the perspective of the users themselves; one of the biographical type⁽²⁹⁾, from an interdisciplinary perspective that integrates cognitive and cultural approaches, which describes and analyzes the ayahuasca treatment processes of four cases of chemical dependence treated at the Amazon Institute of Applied Ethnopsychology (Instituto de Etnopsicologia Aplicada da Amazônia, IDEAA), a center devoted to the treatment of addictions; two in-depth interviews, one that used a qualitative reconstructive method based on the paradigm of Symbolic Interactionism and included interviews with 13 therapists who apply ayahuasca professionally in the treatment of addictions, two specialist researchers, and 14 individuals undergoing ayahuasca-assisted therapy for addictions in diverse contexts in South America⁽²⁶⁾, and the other sought to understand the ritual use of ayahuasca in the treatment and recovery of homeless people by the Flor das Águas Padrinho Sebastião Rescue Unit, in São Paulo⁽¹⁶⁾. Finally, an online retrospective survey, which used 5d-ASC, a psychometrically sound measure of altered states of consciousness (ASCs), to examine ibogaine-induced ASCs, and discusses the demographic characteristics of those seeking ibogaine treatment⁽²²⁾.

With regard to the types of substance use disorder, the studies presented some variations: four focused on tobacco dependence^(17,20,23-24), three on addiction to alcohol^(18-19,21), one to cocaine⁽²⁸⁾ and another six aimed at chemical dependence in general^(16,22,25-27,29). As for the psychotropic drugs used as treatment, five of the studies resorted to psilocybin^(17-20,24), five addressed the use of ayahuasca^(16,26-29), one dealt with ibogaine⁽²²⁾, one addressed the 5-HT2AR agonists⁽²³⁾, one 4-Acetoxy-DMT⁽²⁵⁾, and another referred to the psychedelic experience in general⁽²¹⁾.

In relation to the terms used to define the type of substance consumption, most used the term "addiction"; other terms also found were "substance dependence", "substance use disorder", "adicciones", "alcohol dependence", "tobacco addiction" and "drug dependent state". As for the terms used to refer to the psychedelic agents, most used the nomenclature of the specific substances for each study, although general terms such as "hallucinogens" and "naturalistic psychedelic" were also found. The results of the synthesis of the articles selected are presented in Figure 2.

AUTHOR (YEAR)	COUNTRY	OBJECTIVE	METHODOLOGY	MAIN RESULTS
Gomes et al. (2011) ⁽¹⁶⁾ .	Brazil	To understand the ritual use of ayahuasca in the treatment and recovery of homeless people in São Paulo.	Qualitative research: in- depth interview.	The therapeutic effects of tea are linked to how it is understood and used by the participant.
Johnson et al. (2014) ⁽¹⁷⁾ .	USA	It illustrates a framework for future research studies about the efficacy and mechanisms of hallucinogen-facilitated treatment for addictions.	Open pilot study administering moderate (20 mg/70 kg) and high (30 mg/70 kg) doses of psilocybin.	The smoking cessation rate observed substantially exceeds those commonly reported for other behavioral and/or pharmacological therapies.
Bogenschutz et al. (2018) ⁽¹⁸⁾ .	USA	To illustrate the range of experiences and persistent effects of psilocybin treatment.	Double-blind, placebo- controlled clinical trial of psilocybin-assisted treatment of alcohol use disorder.	Descriptive case studies provide a useful window into the therapeutic process of the psychedelic treatment of addictions.
Nielson et al. (2018) ⁽¹⁹⁾ .	USA	To explore how the patients report on change-related phenomena during post-medication debriefing sessions.	Open pilot study of psilocybin-assisted treatment for alcohol dependence with participants who met the DSM-IV-TR criteria for such dependence.	The participants articulated a series of main phenomena related to changing the outcomes of alcohol consumption and the acute subjective effects of psilocybin.

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AUTHOR (YEAR)	COUNTRY	OBJECTIVE	METHODOLOGY	MAIN RESULTS
Garcia-Romeu et al. (2015) ^{/20)} .	USA	To investigate the potential psychological mechanisms that mediate the results of the psilocybin-facilitated treatment of dependence.	Open pilot study about the treatment of the smoking habit facilitated by psilocybin in the context of the cognitive-behavioral therapy.	The smoking cessation outcomes were significantly correlated with measures of mystical experience on session days.
Garcia-Romeu et al. (2019) ⁽²¹⁾ .	USA	To systematically characterize and determine patterns within cases where the use of psychedelic agents led to self-reported reductions in alcohol misuse.	A cross-sectional study, anonymous from an online survey, with a sample of 343 adults.	Almost all the interviewees reported that they greatly reduced or stopped drinking alcohol since their baseline psychedelic experience.
Heink et al. (2016) ⁽²²⁾ .	USA	It adds to the literature using 5d-ASC, a psychometrically sound measure of altered states of consciousness (ASCs) to examine ibogaine-induced ASCs.	The researchers collected data using an anonymous, online and retrospective survey.	The results indicated a positive correlation between the several dimensions of the ASCs. The participants reported that ibogaine reduced their cravings after treatment, regardless of the ASCs.
Johnson et al. (2017) ⁽²³⁾ .	USA	To complement a laboratory research study on smoking cessation.	An anonymous survey, conducted online with a sample of 358 individuals.	The individual outcomes related to the smoking habit varied, including total abstinence, persistent reductions in the rate of prior smoking, and reductions in regular daily smoking culminating in relapse to the baseline smoking levels.
Johnson et al. (2017) ⁽²⁴⁾ .	USA	To assess the long-term effects of a psilocybin-facilitated smoking cessation program ≥12 months after administration of psilocybin.	It describes biologically verified results of smoking abstinence in the previous pilot study ≥12 months and related data about the subjective effects of psilocybin.	In the 12-month follow-up, 10 participants (67%) were confirmed to be abstinent from smoking. In the long-term follow-up, nine participants (60%) were confirmed to be abstinent from smoking.
Vargas-Perez et al. (2017) ⁽²⁵⁾ .	Canada	The goal is to prove that the 5-HT2A agonists can be used as therapeutic agents to reverse a state of drug dependence, as well as inhibit the aversive effects produced by drug withdrawal.	Administration of non- selective 5-HT2A agonist 4-Acetoxy- Dimethyltryptamine (4-AcO- DMT) in chronic opioids, chronic nicotine or intra-VTA BDNF treated in previously untreated rodents.	Pretreatment with a single injection of systemic 4-AcO- DMT before administration can prevent the rewarding motivational effects of nicotine in addicted mice, blocking withdrawal aversion.
Loizaga-Velder et al. (2014) ⁽²⁶⁾ .	United Kingdom	To explore the ritualistic use of ayahuasca in the treatment of addictions.	An exploratory study using a qualitative reconstructive method based on the paradigm of Symbolic Interactionism.	The results indicate that ayahuasca can serve as a valuable therapeutic tool which, in carefully structured environments, can catalyze neurobiological and psychological processes that support recovery from substance dependence and prevention of relapses.
Talin et al. (2017) ⁽²⁷⁾ .	Without identification	To examine people's healing experiences, that is, their attempts to cure substance use they find problematic.	A qualitative and ethnographically-based analysis of addiction recovery experiences in ayahuasca rituals.	The efficacy of ayahuasca in treating dependence mixes the somatic, symbolic and collective dimensions.
Vitor (2019) ⁽²⁸⁾ .	Brazil	To evaluate the potential of Ayahuasca (AYA) to prevent the expression of behavioral sensitization induced by cocaine and the neurochemical repercussions of the treatment in C57BL/6 mice.	The influence of acute administration of AYA (1.76, 3.0, 17.6 and 30.0 mg/kg of DMT o.r.) on the locomotor activity of the animals (mice) in the open field was evaluated.	Analytical characterization of the drugs and behavioral analysis were performed. The results suggest that AYA can be a good therapeutic strategy for cocaine addiction, paving the way for new studies.
Apud (2019) ⁽²⁸⁾ .	Spain	To describe and analyze the processes of cure of dependence cases treated at the Amazon Institute of Applied Ethnopsychology (<i>Instituto de</i> <i>Etnopsicologia Aplicada da</i> <i>Amazônia</i> , IDEAA).	The procedure involved a qualitative methodology of the biographical type, under an interdisciplinary perspective that integrates cognitive and cultural approaches.	The article concludes by addressing the importance of the ritual's "memory of experience" as a catalyst for new meanings in the patients' biographical narratives.

Figure 2 – Synthesis for the articles selected. Palmas, Tocantins, Brazil, 2020

Discussion

Dependence on alcohol and nicotine are considered chronic disorders that disturb the individuals' psychological and physiological systems, in which they use these substances to generate comfort or alleviate symptoms caused by withdrawal⁽³⁰⁾. Dependence disorders involve a complex neuromodulation process in which these substances alter cellular and molecular aspects of the neurological function, which generate compulsive behavioral effects, but are also influenced by genetic, psychological, social and environmental factors⁽¹⁾.

The pharmacological approach is an important factor in the treatment of alcohol and tobacco dependence, in parallel with the psychological intervention, and the social approach. In general, drugs are used in replacement, detoxification and prevention of relapses⁽¹⁾. The advances in psychopharmacology in the treatment of substance abuse disorders benefited a large number of people; however, due to delay in treatment and to the presentation of several side effects, the pharmacological therapy currently used presents limited effectiveness, as there is low adherence to the use of the medications and/or withdrawal of the recommended therapy⁽³¹⁾.

A promising treatment model is being studied as a complementary therapeutic alternative in a renaissance of research studies on psychedelic substances⁽¹⁷⁻¹⁸⁾. In studies using psilocybin, the mysticomimetic experience can explain some aspects of the positive results of the patient's integration, being a considerable promise in promoting long-term smoking abstinence⁽²⁰⁾.

Other studies indicate that the rituals of the consumer community of ayahuasca, together with social integration, can assist in the treatment of chemical dependence, constituting fluid and adaptable care modalities that play a fundamental role in treatment success, as feeling part of a community has an important therapeutic potential⁽²⁷⁾.

Studies based on experimental and virtual research show self-reported reductions in alcohol and tobacco consumption, some determined by international scores⁽²⁰⁻²¹⁾.

The ritualistic approach with ayahuasca allowed describing the effects in helping to treat dependence. In the first place, the experience is broader than the mere ritualistic process. The patients need to be integrated into the psychedelic experience, to create interactions that produce a memory of this experience; then, the patients' psychedelic experience will be interpreted and re-signified so that, with this, they can establish a link between their new memories, brought by the ritualistic process, and the old memories already existing⁽²⁹⁾.

Studies showing empirical hypotheses with attempts to explain the pharmacodynamics of psychedelic agents were also found^(25-26,28). The partially accepted hypothesis shows the functionalities of ibogaine. Using 5d-ASCs, a psychometrically sound measure of altered states of consciousness (ASCs), it is remarkable that there is a positive correlation between the several dimensions of the ASCs and the patient's improvement, explained by an ability of the ibogaine therapy to provoke changes in the patient's life and desires⁽²²⁾.

Conclusion

The use of psychotropic drugs in the treatment of alcohol and tobacco dependence is not conclusive for the effective treatment of such dependence; however, the use of these entheogenic agents, according to the studies analyzed, shows a window of possibility for an innovation in the current treatments, so that this encompasses the patients' psychedelic experience, their interaction and co-participation in the treatment.

It was evident in the studies addressed that the experience itself, provided by the psychedelic agents, is broad, and that its therapeutic management needs to include integration activities that contribute to treatment innovation. Another important element is the creation of a memory of the experience with psychedelic agents, which is interpreted and re-signified, so that, from a biographical point of view, it becomes the key articulation point between an old and a new personality.

It was also observed that mystical-mimetic experiences are perceived as a relevant contribution to the treatments. Some patients report, and scientific scales confirm, very intense individual experiences similar to mystical states, which produce a sense of well-being and personal growth.

This study sought to collaborate with the systematization of knowledge on the theme, stimulating further discussions on the use of psychedelic agents in the treatment of alcohol and tobacco dependence, in order to help expand treatment perspectives, as well as show new contributions to the therapeutic methods.

References

 Silva MTB, Araujo FLO, Félix FHC, Simão AFL, Lobato RFG, Sousa FCF, et al. Alcohol and nicotine. Rev Neuroci. 2010;18(4):531-7. https://doi.org/10.34024/rnc.2010. v18.8436

2. World Health Organization (WHO). Global status report on alcohol and health [Internet]. Geneva: WHO; 2018 [cited 2020 Dec 21]. Available from: https://apps.who. int/iris/handle/10665/274603

3. Fundação Oswaldo Cruz (BR), Instituto de Comunicação e Informação Científica e Tecnológica em Saúde. III Levantamento Nacional sobre o uso de drogas pela população brasileira [Internet]. Rio de Janeiro: FIOCRUZ/ ICICT; 2017 [cited 2021 Aug 10]. 528 p. Available from: https://www.arca.fiocruz.br/bitstream/icict/34614/1/ III%20LNUD_PORTUGU%C3%8AS.pdf 4. Organização Mundial da Saúde (OMS). Transtornos devido ao uso de substâncias. Relatório sobre a saúde no mundo. Saúde Mental: nova concepção, nova esperança [Internet]. 2001 [cited 2021 Aug 10]. Available from: https://www.dgs.pt/documentos-e-publicacoes/relatorio-mundial-da-saude-2001--saude-mental-nova-concepcao-nova-esperanca-pdf.aspx

8

5. Frossard SC. As Políticas Públicas e as Comunidades Terapêuticas nos Atendimentos à Dependência Química. Serv Soc Rev [Internet]. 2009 [cited 2021 Aug 10];11:1-14. Available from: http://www.uel.br/revistas/ssrevista/ pdf/2009/29%20AS%20POL%CDCAS%20P%DABLICAS%20 E%20AS%20COMUNIDADE%20TERAP%CAUTICAS-COM%20REVIS%C30%20D0%20AUTOR.pdf

6. Pinto M, Ugá MA. The cost of tobacco-related diseases for Brazil's Unified National Health System. Cad Saúde Pública. 2010;26(6):1234-45. https://doi.org/10.1590/ S0102-311X2010000600016

7. Organização Mundial de Saúde (OMS). Neurociência do uso e da dependência de substâncias psicoativas [Internet]. São Paulo: Roca; 2006 [cited 2021 Aug 10]. Available from: http://apps.who.int/iris/ bitstream/handle/10665/42666/9788572416665_por. pdf?sequence=2

8. Coutinho ESF, Bahia L, Barufaldi LA, Abreu GA, Malhão TA, Pepe CR, et al. Cost of diseases related to alcohol consumption in the Brazilian Unified Health System. Rev Saúde Pública. 2016;50. https://doi.org/10.1590/ S1518-8787.2016050005741

Organização Pan-Americana da Saúde (OPAS);
Organização Mundial da Saúde (OMS). Álcool [Internet].
2020 [cited 2021 Aug 10]. Available from: https://www.paho.org/pt/topicos/alcool

10. Souza LG, Menandro M, Menandro P. Alcoholism, its causes and treatment in the social representations constructed by Brazilian Family Health professionals. Physis. 2015;25(4):1335-60. https://doi.org/10.1590/ S0103-73312015000400015

11. Ministério da Saúde (BR), Secretaria de Ciência, Tecnologia e Insumos Estratégicos, Departamento de Gestão e Incorporação de Tecnologias em Saúde. Protocolo Clínico e Diretrizes Terapêuticas do Tabagismo [Internet]. Brasília: Ministério da Saúde; 2019 [cited 2021 Aug 10]. Available from: http://conitec.gov.br/images/ Relatorios/2020/Relatrio_PCDT_Tabagismo_520_2020_ FINAL.pdf

12. Labate BC, Santos RG, Anderson B, Mercante M, Barbosa PCR. Considerações sobre o tratamento da dependência por meio da ayahuasca. [s.l.]: Núcleo de Estudos Interdisciplinares sobre Psicoativos; 2009 [cited 2021 Aug 10]. Available from: http://neip.info/novo/ wp-content/uploads/2015/04/labate_et_all_tratamento_ dependencia_ayahuasca_final.pdf Johnson MW, Griffiths RR. Potential Therapeutic Effects of Psilocybin. Neurotherapeutics. 2017;14:734 40. https://doi.org/10.1007/s13311-017-0542-y

14. Souza MT, Silva MD, Carvalho R. Integrative review: what is it? How to do it? Einstein (São Paulo). 2010;8(1):102-6. https://doi.org/10.1590/ s1679-45082010rw1134

15. Mendes KDS, Silveira RCCP, Galvão CM. Integrative literature review: a research method to incorporate evidence in health care and nursing. Texto Contexto Enferm. 2008;17(4):758-64. https://doi.org/10.1590/ S0104-07072008000400018

16. Gomes BR. The meaning of ritualized use of ayahuasca in the treatment of homeless users in a group in São Paulo [Master's thesis]. São Paulo: Faculdade de Saúde Pública, Universidade de São Paulo; 2011. https://doi. org/10.11606/D.6.2011.tde-09092011-160114

17. Johnson MW, Garcia-Romeu A, Cosimano MP, Griffiths RR. Pilot study of the 5-HT2AR agonist psilocybin in the treatment of tobacco addiction. J Psychopharmacol. 2014;28(11):983-92. https://doi. org/10.1177/0269881114548296

18. Bogenschutz MP, Podrebarac SK, Duane JH, Amegadzie SS, Malone TC, Owens LT, et al. Clinical Interpretations of Patient Experience in a Trial of Psilocybin-Assisted Psychotherapy for Alcohol Use Disorder. Front Pharmacol. 2018;9:100. https://doi.org/10.3389/fphar.2018.00100 19. Nielson EM, May DG, Forcehimes AA, Bogenschutz MP. The Psychedelic Debriefing in Alcohol Dependence Treatment: Illustrating Key Change Phenomena through Qualitative Content Analysis of Clinical Sessions. Front Pharmacol. 2018;9:132. https://doi.org/10.3389/ fphar.2018.00132

20. Garcia-Romeu A, Griffitts R, Johnson M. Psilocybin-Occasioned Mystical Experiences in the Treatment of Tobacco Addiction. Curr Drug Abuse Rev. 2015;7(3):157-64. https://doi.org/10.2174/18744737086661501071 21331

21. Garcia-Romeu A, Davis AK, Erowid F, Erowid E, Griffiths RR, Johnson MW. Cessation and reduction in alcohol consumption and misuse after psychedelic use. J Psychopharmacol. 2019;33(9):1088-101. https://doi. org/10.1177/0269881119845793

22. Heink A, Katsikas S, Lange-Altman T. Examination of the Phenomenology of the Ibogaine Treatment Experience: role of altered states of consciousness and psychedelic experiences. J Psychoactive Drugs. 2017;49(3):201-8. https://doi.org/10.1080/02791072.2017.1290855

23. Johnson MW, Garcia-Romeu A, Johnson PS, Griffiths RR. An online survey of tobacco smoking cessation associated with naturalistic psychedelic use. J Psychopharmacol. 2017;31(7):841-50. https://doi. org/10.1177/0269881116684335

24. Johnson MW, Garcia-Romeu A, Johnson PS, Griffiths RR. Long-term follow-up of psilocybin-facilitated smoking cessation. Am J Drug Alcohol Abuse. 2017;43:1-6. https://doi.org/10.3109/00952990.2016.1170135

25. Vargas-Perez H, Grieder TE, Ting-A-Kee R, Maal-Bared G, Chwalek M, Kooy D. A single administration of the hallucinogen, 4-acetoxy-dimethyltryptamine, prevents the shift to a drug-dependent state and the expression of withdrawal aversions in rodents. Eur J Neurosci. 2017;45(11):1410-7. https://doi.org/10.1111/ejn.13572 26. Loizaga-Velder A, Verres R. Therapeutic Effects of Ritual Ayahuasca Use in the Treatment of Substance Dependence-Qualitative Results. J Psychoactive Drugs. 2014;46(1):63-72. https://doi.org/10.1080/02791072.2013.873157

27. Talin P, Sanabria E. Ayahuasca's entwined efficacy: an ethnographic study of ritual healing from addiction. Int J Drug Policy. 2017;44(1):23-30. https://doi.org/10.1016/j. drugpo.2017.02.017

28. Bruno V. Ayahuasca in the treatment of cocaine addiction: possible therapeutic interventions? [Master's thesis]. São Paulo: Faculdade de Ciências Farmacêuticas, Universidade de São Paulo; 2019. https://doi. org/10.11606/D.9.2019.tde-09122019-185108

29. Apud I. Ayahuasca in the treatment of addictions. Study of four cases treated in IDEAA, using an interdisciplinary perspective. Interdisciplinaria: Rev Psicol y Ciencias Afines. 2019;36(1):1-7. https://doi. org/10.16888/interd.2019.36.1.10

30. Swift RM, Lewis DC. Farmacologia da Dependência e Abuso de Drogas. In: Armstrong AW, Tashjian AH, Golan DE. Princípios De Farmacologia: A Base Fisiopatologica da Farmacoterapia [Internet]. 2. ed. Rio de Janeiro: Guanabara Koogan; 2009 [cited 2021 Aug 10]. p. 260-78. Available from: https://edisciplinas.usp.br/pluginfile. php/3319241/mod_resource/content/1/Farmacologia%20 da%20dependencia%20e%20abuso%20%20de%20 drogas.pdf

31. Tang GY, Parekh J. Side effects of addiction treatment. Science. 2018;361(6404):761. https://doi.org/10.1126/ science.aau6548

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